REMARKS

The present amendment is being submitted at the Examiner's request since the amendment of August 20, 2007 incorrectly amended pages 41 and page 42. The present amendment correctly amends pages 41 and 42 and a marked-up copy of these pages is submitted with the handwritten pages so the amendment can be seen properly.

Respectfully submitted,

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CAM:mlp Enclosures 10

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$$A - (CH_2)_{\overline{m}} + R_{36} + V - \Phi - (XXII)$$

$$(XXII) \qquad (XXII)$$

Diagram 12

The amines of general formula (V), Diagram 13, in which A, X, R₃₉, R₃₇, p and q are as defined above, can also be prepared by condensation of an amine of general formula (XXIII) with a commercial aldehyde of general formula (XXIV) in a reducing medium under the conditions described previously. The secondary amines of general formula (V) are then protected in the form of *tert*-butyl carbamate, under the conditions described previously. The syntheses of the amines of general formula (XXIII) are described in the "Preparation of the Intermediates" chapter. Moreover, the non-commercial aldehydes of general formula (XXVI) can be prepared according to *J. Org. Chem.*, 1993, 58, 1385-92.

$$A-X-(CH_2) = N \begin{pmatrix} R_{37} \\ H \end{pmatrix} + \begin{pmatrix} CH_2 \\ Q-1 \end{pmatrix} = \begin{pmatrix} R_{39} \\ NO_2 \end{pmatrix} + \begin{pmatrix} R_{39} \\ (Boc)_2 \end{pmatrix}$$
(XXIII) (XXIV)

Diagram_13

Synthesis of the amines by reduction of the carboxamides:

The amines of general formula (V), Diagram 14, in which A, X, R₃₉, R₃₇ and q are as defined above, are accessible by reduction of the carboxamide derivatives of general formula (V), the synthesis of which is described in the "Synthesis of the carboxamides" chapter (Diagram 4). The reduction stage is carried out in an anhydrous medium, by heating at 70-80 °C, in the presence of a carboxamide selective reagent such as, for example, BH₃. THF, in a solvent such as, for example, THF. The secondary amines thus prepared can be protected in the form of *tert*-butyl carbamate under the conditions described previously.

<u>Diagram 14</u>

Synthesis of the ureas:

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The ureas of general formula (V), Diagram 15, in which A, Y, Φ , R₃₉, R₃₆ and R₃₇ are as defined above, are accessible by reaction of the amines of general formula (XIX) with the amines of general formula (XXII) in the presence of triphosgene and of a base such as, for example, diisopropylethylamine in an inert solvent such as dichloromethane according to an experimental protocol described in *J. Org. Chem.* (1994) 59 (7), 1937-1938.

$$A-N$$
 H
 $+$
 R_{37}
 $-Y-\Phi$
 NO_2
 NO_2
 $(XXII)$

Diagram 15

10 Synthesis of pyrrolidinyl-pyrimidine derivatives:

The pyrimidine derivatives of general formula (V), Diagram 16, in which Q represents piperazine, Φ and R₃₉ being as defined above, are prepared by condensation of 4-chloro-2,6-di-pyrrolidin-1-yl-pyrimidine (*J. Med. Chem.* (1990) 33 (4), 1145-1151) with the piperazine derivatives of general formula (X), by heating, for 24 to 48 hours, in anhydrous pyridine at a temperature of 80-110 °C.

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